5

10

ABSTRACT OF THE INVENTION

The present invention relates to compositions comprising proteasome inhibitors, such as lactocystin and analogues thereof. These compositions are used for the following purposes: (1) to disrupt mitochondrial function (useful against cancer, inflammation, adverse immune reaction and hyperthyroidism), (2) to disrupt nitric oxide synthesis (useful against inflammation and septic shock), and (3) to reverse ongoing adverse immune reactions, such as autoimmune diseases and graft rejection. In the later case, the compositions are administered once the patients' T cells are mostly activated. Proteasome inhibitors can also be combined to immuno-suppressive drugs like rapamycin, cyclosporin A and FK506. Finally, a method for screening a compound having a proteasome inhibition activity is also disclosed and claimed.

Express Mail" mailing label number EL353244453US
Express Mail" mailing label number
Date of Denosit June 29, 1999
heraby certify that this paper or fee is being deposited
with the United States Postal Service "Express Mail Post
Office to Addressee" service under 37 CFR 1.10 on the
date indicated above and is addressed to the Commis-
sioner of Patents and Trademarks, Washington, D.C. 20231
CIONS WAS
χ printed name
Dun the
Signature